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(54) Title: GENES AND PROTEINS, AND THEIR USE (57) Abstract		Published <i>Without international search report and to be republished upon receipt of that report.</i>																																																			
<p>According to the present invention, a series of genes are identified in Group B <i>Streptococcus</i>, the products of which may be associated with the outer surface of the organism. The genes, or functional fragments thereof, may be useful in the preparation of therapeutics, e.g. vaccines to immunise a patient against microbial infection.</p>																																																					

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GENES AND PROTEINS, AND THEIR USEField of the Invention

This invention relates to the identification of
5 bacterial genes and proteins, and their use. More particularly, it relates to their use in therapy, for immunisation and in screening for drugs.

Background to the Invention

Group B *Streptococcus* (GBS), also known as
10 *Streptococcus agalactiae*, is the causative agent of various conditions. In particular, GBS causes:

Early onset neonatal infection.

This infection usually begins in utero and causes severe septicaemia and pneumonia in infants, which is
15 lethal if untreated and even with treatment is associated with a 10-20% mortality rate.

Late onset neonatal infection.

This infection occurs in the period shortly after birth until about 3 months of age. It causes a
20 septicaemia, which is complicated by meningitis in 90% of cases. Other focal infections also occur including osteomyelitis, septic arthritis, abscesses and endophthalmitis.

Adult infections.

25 These appear to be increasingly common and occur most frequently in women who have just delivered a baby, the elderly and the immunocompromised. They are characterised by septicaemia and focal infections including osteomyelitis, septic arthritis, abscesses and
30 endophthalmitis.

Urinary tract infections.

GBS is a cause of urinary tract infections and in pregnancy accounts for about 10% of all infections.

Veterinary infections.

35 GBS causes chronic mastitis in cows. This, in turn, leads to reduced milk production and is therefore of considerable economic importance.

GBS infections can be treated with antibiotics. However, immunisation is preferable. It is therefore desirable to develop an immunogen that could be used in a therapeutically-effective vaccine.

5 Summary of the Invention

The present invention is based on the identification of a series of genes in GBS, and also related organisms, the products of which may be localised on the outer surface of the organism and therefore may be used as a target for
10 immuno-therapy.

According to one aspect of the invention, a peptide is encoded by an operon including any of the genes identified herein as pho1-13, pho3-21, pho2-15, pho3-18, pho3-22, pho3-3, pho3-17, pho2-2, pho1-5, pho3-1, pho3-23, pho3-50,
15 pho1-14, pho2-10, pho3-14, pho3-24 and pho3-29, obtainable from Group B *Streptococcus*, or a homologue or functional fragment thereof. Such a peptide is suitable for therapeutic use, e.g. when isolated.

The term "functional fragments" is used herein to
20 define a part of the gene or peptide which retains the activity of the whole gene or peptide. For example, a functional fragment of the peptide may be used as an antigenic determinant, useful in a vaccine or in the production of antibodies.

25 A gene fragment may be used to encode the active peptide. Alternatively, the gene fragment may have utility in gene therapy, targetting the wild-type gene *in vivo* to exert a therapeutic effect.

A peptide according to the present invention may
30 comprise any of the amino acid sequences identified herein as SEQ ID NOS. 2, 4, 6, 8, 10, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33 and 35, or a functional fragment thereof.

Because of the extracellular or cell surface location,
35 the peptides of the present invention may be suitable candidates for the production of therapeutically-effective vaccines against GBS. The term "therapeutically-effective"

is intended to include the prophylactic effect of vaccines. For example, a vaccine may comprise a peptide according to the invention, or the means for its expression, for the treatment of infection. The vaccine may be administered to females prior to or during pregnancy to protect mother and neonate against infection by GBS.

According to another aspect of the invention, the peptides or genes may be used for screening potential antimicrobial drugs or for the detection of virulence.

A further aspect of this invention is the use of any of the products identified herein, for the treatment or prevention of a condition associated with infection by a Group B Streptococcal strain.

Although the protein has been described for use in the treatment of patients, veterinary uses of the products of the invention are also considered to be within the scope of the present invention. In particular, the peptides or the vaccines may be used in the treatment of chronic mastitis, especially in cows.

Description of the Invention

The present invention is described with reference to Group B Streptococcal strain M732. However, all the GBS strains and many other bacterial strains are likely to include related peptides or proteins having amino acid sequence homology with the peptide of M732. Organisms likely to contain the peptides include, but are not limited to, *S. pneumoniae*, *S. pyogenes*, *S. suis*, *S. milleri*, Group C and Group G Streptococci and Enterococci. Vaccines to each of these may be developed in the same way as described for GBS.

Preferably, the peptides that may be useful for the production of vaccines have greater than 40% sequence similarity with the peptides identified herein. More preferably, the peptides have greater than 60% sequence similarity. Most preferably, the peptides have greater than 80% sequence similarity, e.g. 95% similarity.

Having characterised a gene according to the invention, it is possible to use the gene sequence to establish homologies in other microorganisms. In this way it is possible to determine whether other microorganisms have similar outer surface products. Sequence homologies may be established by searching in existing databases, e.g. EMBL or Genbank.

Peptides or proteins according to the invention may be purified and isolated by methods known in the art. In particular, having identified the gene sequence, it will be possible to use recombinant techniques to express the genes in a suitable host. Active fragments and homologues can be identified and may be useful in therapy. For example, the peptides or their active fragments may be used as antigenic determinants in a vaccine, to elicit an immune response. They may also be used in the preparation of antibodies, for passive immunisation, or diagnostic applications. Suitable antibodies include monoclonal antibodies, or fragments thereof, including single chain fv fragments. Methods for the preparation of antibodies will be apparent to those skilled in the art.

The preparation of vaccines based on attenuated microorganisms is known to those skilled in the art. Vaccine compositions can be formulated with suitable carriers or adjuvants, e.g. alum, as necessary or desired, and used in therapy, to provide effective immunisation against Group B Streptococci or other related microorganisms. The preparation of vaccine formulations will be apparent to the skilled person.

More generally, and as is well known to those skilled in the art, a suitable amount of an active component of the invention can be selected, for therapeutic use, as can suitable carriers or excipients, and routes of administration. These factors will be chosen or determined according to known criteria such as the nature/severity of the condition to be treated, the type or health of the subject etc.

The products of the present invention were identified as follows:

A partial gene library of GBS (strain M732) chromosomal DNA was prepared using the plasmid vectors pFW-
5 phoA1, pFW-phoA2 and pFW-phoA3 (Podbielski, A. et al. 1996. Gene 177:137-147). These plasmids possess a constitutive spectinomycin adenyltransferase antibiotic resistance marker, which confers a high level of spectinomycin resistance and is therefore easily selected. Furthermore,
10 these vectors contain a truncated (leaderless) *Escherichia coli* phoA gene for alkaline phosphatase. The three vectors differ only with respect to the reading frame in which the leaderless phoA gene exists, as compared to an upstream in-frame BamHI restriction enzyme site. Because this
15 truncated *E. coli* phoA gene lacks the appropriate leader sequence for export of this enzyme across the bacterial membrane, extracellular alkaline phosphatase activity is absent when these plasmids are propagated in an *E. coli* phoA mutant (e.g. strain DH5 α). The chromogenic alkaline
20 phosphatase substrate, XP (5-bromo-4-chloro-3-indolyl-phosphate), does not enter intact bacterial cells and therefore only exported or surface associated alkaline phosphatase activity can be detected. When exported or surface associated alkaline phosphatase activity is
25 present, the chromogenic XP substrate is cleaved to yield a blue pigment and the corresponding bacterial colonies can be identified by their blue colour.

Plasmid DNA was digested to completion with BamHI and dephosphorylated using shrimp alkaline phosphatase. GBS
30 genomic DNA was partially digested with Sau3AI, size fractionated on a sucrose gradient and fragments <1kb in size were ligated into the prepared pFW-phoA vectors. *E. coli* strain DH5 α was chosen as the cloning host since it lacks a functional phoA gene. Recombinant plasmids were
35 selected on Luria agar containing 100 μ g/ml of spectinomycin and 40 μ g/ml of the chromogenic XP substrate. *E. coli* transformants harbouring plasmids containing GBS

insert DNA that complements the export signal sequence of the leaderless *phoA* gene were identified by the blue colour of the colonies. Approximately 30000 different recombinant plasmids containing GBS insert DNA were screened in this manner and 83 recombinant plasmids, which complemented the leaderless *phoA*, were chosen for further study.

From these experiments, several clones were selected each containing a plasmid containing a gene (or part thereof), which complemented the leaderless *phoA*.

Having identified the gene in each clone it is then possible to obtain the full-length gene sequence, as follows.

Using the identified and sequenced gene fragment, oligonucleotide primers were designed for genomic DNA sequencing. These primers were designed so as to sequence in an 'outward' direction from the obtained sequence. Once read, the sequence obtained was checked to see if the 5' and 3' termini of the gene had been reached. The presence of these features was identified by checking against homologous sequences, and for the 5' end the presence of an AUG start codon (or accepted equivalent) preceded by a Shine-Dalgarno consensus sequence, and for the 3' end, the presence of a translation termination (Stop) codon.

Upon identification of the full-length gene, primers were designed for amplification of full-length product. Primers used included restriction enzyme recognition sites (NcoI at the 5' end and EcoO109I at the 3' end) to allow subsequent cloning of the product into the Lactococcal expression system used.

PCR was carried out using the primers, and the products cloned into a PCR 2.1 cloning vector (In Vitrogen). Following confirmation of the presence of the cloned fragment, the DNA was excised using the restriction enzymes NcoI and EcoO109I.

The vector into which this fragment was inserted was a modified version of pNZ8048 (Kuipers, O. P. et al. (1998) J. Biotech 64: 15-21). This vector, harbouring a

lactococcal origin of replication, a chloramphenicol resistance marker, an inducible nisin promoter and a multicloning site was altered by the replacement of the multicloning site with two 10X His tags, flanked on the 5-
5 most end with an NcoI site, split in the middle with a multicloning site (including an Eco0109I site), and a Stop (termination) codon at the 3'end of the His tags.

The gene of interest was inserted so that a 10X His tag was in the 3' position relative to the coding region.
10 Following transformation of the recombinant plasmid into *L.lactis* (strain NZ9000 - Kuipers, O. P. et al. (1998) *supra*), a 400 ml liquid culture was set up and translation of the protein was induced by the addition of nisin to the culture. After a 2 hour incubation, the cells were
15 harvested and lysed by bead beating. The resultant lysate was cleared by centrifugation, then passed over a metal affinity (Talon, Clontech) column. The column was washed repeatedly before bound proteins were eluted with Imidazole.

20 To identify fractions containing the His-tagged recombinant protein, an aliquot from each fraction was analysed by SDS-PAGE, Western blotted and probed with anti-His antibodies.

The recombinant protein obtained was then used to
25 immunise New Zealand white rabbits, with pre-immune sera being harvested prior to immunisation. Following a boost, the rabbits were sacrificed and sera collected. This sera was used in Western blots, ELISA and animal protection models.

30 Using the sera obtained from the animal studies, immunosorption studies were carried out.

Group B *Streptococcus* was grown in 20ml Todd Hewitt broth (THB) for 8 hours, harvested and resuspended in 5ml PBS. 50µl aliquots of this were used to coat wells in a 96
35 well plate (Nunc Immuno-Sorb). This was left at 4°C overnight to allow for adsorbance of the bacteria onto the plate. Plates were washed twice with PBS, then blocked

with 3%BSA in PBS for 1hr at 37°C. Plates were again washed. Serial 10 fold dilutions of the sera were made in PBS and 50µl of these dilutions were added to the wells of the plate, in duplicate. The plate was covered and
5 incubated for 1 hr at 37°C. The plate was washed, then 50µl anti-rabbit alkaline phosphatase conjugated secondary antibody at a concentration of 1:5000 was added to each well. Following incubation at 37°C for an hour, the plate was washed again. 50µl substrate (PNPP) was added to each
10 well, and the reaction allowed to proceed for 30min before the adsorbance was read at 405 nm.

Animal protection studies were also carried out to test the effectiveness of protection on the immunised rabbits.

15 GBS M732 was grown up in THB until mid-log phase was reached - approximately 5 hours. Cells were counted in a counting chamber, and bacteria were diluted to give a concentration of 2×10^7 bacteria per ml in pre-immune or test sera. 50µl of this was injected via the
20 intraperitoneal route into 0-1 day old mice. The mice were observed for survival over 48 hours.

The following Examples illustrate the invention.

Example 1

25 A first clone contained a gene sequence identified herein as SEQ ID NO. 1, with an amino acid sequence identified as SEQ ID NO. 2, and classified as pho1-13.

A comparison of the amino acid sequence of pho1-13 was performed.

30 Homologues to the GBS pho1-13 gene product can be identified in *Streptococcus pyogenes*, *S. pneumoniae*, *S. salivarius*, *Escherichia coli*, *Yersinia enterocolitica*, *Aquifex aeolicus*, *Helicobacter pylori* and *Haemophilus influenzae*. The *S. pyogenes* and *S. pneumoniae* homologues
35 were identified from genome sequence data and no annotations were available as to the identity of the gene or gene products. In all other cases, the above homologues

can be identified as ATP-dependent Clp protease proteolytic subunits. The catalytic activity of Clp proteases results in the hydrolysis of proteins to small peptides in the presence of ATP and magnesium (Giffard, P.M. et al. 1993. J. Gen. Microbiol. 139:913-920). Furthermore, the ClpP component of Clp proteases has been shown to be induced as part of the heat shock response (Kroh, H.E. and L.D. Simon. 1990. J. Bacteriol. 172:6026-6034) and it is probable that this subunit or the complete proteolytic domain would associated with the bacterial surface.

Immunisation studies, carried out as described above, yielded the following results.

Treatment	No animals	No animals surviving	
		at time (hrs)	
		24	48
PBS	10	7	0
Pre-immunised	37	13	0
Immunised	38	17	9

Example 2

A second clone was selected containing a plasmid designated pho1-14. This plasmid contained a gene (or part thereof), which complemented the leaderless *phoA*. The nucleotide and deduced amino acid sequences are shown as SEQ ID NOS. 3 and 4, respectively.

A comparison of the amino acid sequence of pho1-14 was performed.

Homologues to the GBS pho1-14 gene product can be identified in *Streptococcus pyogenes*, *Enterococcus faecalis* and *Streptococcus pneumoniae*. These homologues were identified from genome sequence data and no annotations were available as to the identity of the gene or gene products. Additionally, two possible homologues were also

identified from *Shigella flexneri* (SpaR) and *Yersinia pseudotuberculosis* (YscT). These latter two homologues are related proteins, believed to be anchored in the bacterial membrane (Bergman, T. et al. 1994. J. Bacteriol. 176:2619-
5 2626). In *S. flexneri*, the product of the *spaR* gene has been shown to be important for invasion of epithelial cells (Sasakawa, C. et al. 1993. J. Bacteriol. 175:2334-2346). Furthermore, the product of the *spaR* gene is also required for surface presentation of invasion plasmid antigens. The
10 analogous protein in *Y. pseudotuberculosis* is a component of the Yop secretion system and is also important for virulence in this organism.

Example 3

A third clone was selected containing a plasmid
15 designated pho1-5. This plasmid contained a gene (or part thereof), which complemented the leaderless *phoA*. The nucleotide and deduced amino acid sequences are shown as SEQ ID NOS. 5 and 6.

A comparison of the amino acid sequence of pho1-5 was
20 performed.

Homologues to the GBS pho1-5 gene product can only be identified in *Streptococcus pyogenes* and *Staphylococcus carnosus* (*sceA*). The *S. pyogenes* homologue was identified from genome sequence data and no annotations were available
25 as to the identity of the gene or gene products. Furthermore, little information is available on the function of the *sceA* gene product from *S. carnosus*. The *sceA* gene product shows some sequence similarity to the aggregation promoting protein from *Lactobacillus gasseri*.
30 Based on analysis of the *sceA* gene product, this molecule contains a well-conserved signal sequence and is apparently secreted or associated with the bacterial cell surface.

Example 4

A further clone was selected containing a plasmid
35 designated pho3-3. This plasmid contained a gene (or part thereof), which complemented the leaderless *phoA*. The

nucleotide and deduced amino acid sequences are shown as SEQ ID NOS. 7 and 8.

A comparison of the amino acid sequence of pho3-3 was performed.

5 Homologues to the GBS pho3-3 gene product can be identified in *Streptococcus mutans* (rmlC), (cpsM) *S. pneumoniae* and *S. pyogenes*. The *S. pyogenes* homologue was identified from genome sequence data and no annotations were available as to the identity of the gene or gene
10 product. In *S. pneumoniae*, the homologue can be identified as dTDP-4-keto-6-deoxy glucose-3,5-epimerase. In the other two cases, the above homologues can be identified as dTDP-4-keto-L-rhamnose reductase (rmlC). In *S. mutans*, the gene encoding this enzyme, *rmlC*, is part of the *rml* locus.
15 The *rml* locus consists of three genes which exhibit significant similarity to enzymes involved in the biosynthesis of dTDP-rhamnose, the immediate precursor of the rhamnose component in the *S. mutans* polysaccharide capsule (Tsukioka, Y. et al. 1997. J. Bacteriol. 179:1126-
20 1134). An analogous locus has also been identified in *S. pneumoniae* (Coffey, T.J. et al. 1998. Mol. Microbiol. 17:73-83). Almost all *Streptococci* characteristically possess rhamnose in their cell wall associated polysaccharides (Schleifer, K.H. and R. Kilper-Bälz. 1987. Syst. Appl. Microbiol. 10:1-19), and it is highly probable that dTDP-4-keto-L-rhamnose reductase would be associated with the outer surface in *Streptococci*.

Example 5

A further clone was selected containing a plasmid
30 designated pho2-10. This plasmid contained a gene (or part thereof), which complemented the leaderless *phoA*.

The nucleotide sequence is shown as SEQ ID NO. 9. From this, upstream and downstream coding regions were identified, and the deduced amino acid sequences shown as
35 SEQ ID NOS. 10 and 11.

A comparison of the amino acid sequences of pho2-10 was performed.

Homologues to the GBS pho2-10 gene product can be identified in *Streptococcus pyogenes*, *Enterococcus faecalis*, *Debaryomyces occidentalis* (hatI) and *Escherichia coli* (trkD). The *S. pyogenes* and *E. faecalis* homologues were identified from genome sequence data and no annotations were available as to the identity of the gene or gene products. In the yeast *D. occidentalis*, the *hak1* gene is a homologue of the *trkD* gene from *E. coli* (Banuelos, M.A. et al. 1995. EMBO J. 14:3021-3027). The *trkD* gene of *E. coli* is part of the *kup* potassium uptake system. The specific homolog identified here is the *kup* system potassium uptake protein. The *kup* system is a constitutive potassium uptake system in *E. coli*. The *kup* system potassium uptake protein contains a highly hydrophobic N-terminus that is predicted to span the membrane at least 12 times. Kup is not homologous to other known membrane protein sequences. There is no indication of ATP binding, and it is proposed that the system is driven by a chemiosmotic gradient (Schleyer, M. & E.P. Bakker, 1993. J. Bacteriol. 175:6925-6931).

Example 6

A further clone was selected containing a plasmid designated pho2-15. This plasmid contained a gene (or part thereof), which complemented the leaderless *phoA*. The nucleotide and deduced amino acid sequences of the gene are shown as SEQ ID NOS. 12 and 13.

A comparison of the amino acid sequence of pho2-15 was performed.

Homologues to the GBS pho2-15 gene product can be identified in *Streptococcus pyogenes*, *Streptococcus pneumoniae*, *Enterococcus faecalis* and *Escherichia coli* (*gatC* and *SgCC*). The *S. pyogenes*, *S. pneumoniae* and *E. faecalis* homologues were identified from genome sequence data and no annotations were available as to the identity of the gene or gene products. In *E. coli*, the *gatC* and *sgCC* gene products can be identified as being the IIC component of phosphoenolpyruvate-dependent sugar

phosphotransferase systems (PTS), a major carbohydrate active-transport system. In PTS systems, the IIC component is typically involved in binding of extracellular carbohydrates and forms a complex with the IID component to constitute a membrane channel (Nobelmann, B. and J.W. Lengeler. 1995. Biochim. Biophys. Acta 1262:69-72).

Example 7

A further clone was selected containing a plasmid designated pho2-2. This plasmid contained a gene (or part thereof), which complemented the leaderless *phoA*. The nucleotide and deduced amino acid sequences of the gene are shown as SEQ ID NOS. 14 and 15, respectively.

A comparison of the amino acid sequence of pho2-2 was performed.

Homologues to the GBS pho2-2 gene product can be identified in *Enterococcus faecalis*, *Escherichia coli* (malK and afuC), *Bacillus subtilis* (glnO), *Haemophilus influenzae* (yebM and potA), *Streptococcus pyogenes*, *Streptococcus pneumoniae* and *Salmonella typhimurium* (malK). The *E. faecalis*, *S. pyogenes* and *S. pneumoniae* homologues were identified from genome sequence data and no annotations were available as to the identity of the gene or gene products. In all other cases, homologues represented ATP-binding transport proteins that are part of ABC type transporters. Many of the components of ABC type transporters are membrane or cell surface associated, as these systems are involved in the transport of macromolecules from the extracellular environment to the intracellular compartment.

Example 8

A further clone was selected containing a plasmid designated pho3-14. This plasmid contained a gene (or part thereof), which complemented the leaderless *phoA*. The nucleotide and deduced amino acid sequences of the gene are shown as SEQ ID NOS. 16 and 17.

A comparison of the amino acid sequence of pho3-14 was performed and no homologues could be identified in any of

the public databases. One homologue to the GBS pho3-14 gene product can be identified in *Streptococcus pyogenes*, but this homologue was identified from genome sequence data and no annotations were available as to the identity of the gene or gene product. Using this *S. pyogenes* homologue to search the public databases yielded no further information. Since the pho3-14 product complemented the leaderless *phoA* gene, it can be concluded that this protein (or part thereof) would most probably be located extracellularly.

10 Example 9

A further clone was selected containing a plasmid designated pho3-17. This plasmid contained a gene (or part thereof), which complemented the leaderless *phoA*. The nucleotide and deduced amino acid sequences of the gene are shown as SEQ ID NOS. 18 and 19.

A comparison of the amino acid sequence of pho3-17 was performed.

Homologues to the GBS Pho3-17 gene product can be identified in *Streptococcus mutans* and *Lactococcus lactis*, with similarity being shown to N-acetyl muramidase. Similarity is also seen with an unidentified gene, yubE from *Bacillus subtilis*.

N-acetylmuramidase is an autolysin that is involved in cell division. Using this limited information along with the fact that pho3-17 complemented the leaderless *phoA* gene, it can be concluded that the pho3-17 product would most probably be located extracellularly.

Example 10

A further clone was selected containing a plasmid designated pho3-18. This plasmid contained a gene (or part thereof), which complemented the leaderless *phoA*. The nucleotide and deduced amino acid sequences of the gene are shown as SEQ ID NOS. 20 and 21.

A comparison of the amino acid sequence of pho3-18 was performed.

Homologues to the GBS pho3-18 gene product can be identified in *Streptococcus pyogenes* and *Streptococcus*

pneumoniae. These homologues were identified from genome sequence data and no annotations were available as to the identity of the gene or gene products. Using these *S. pyogenes* and *S. pneumoniae* homologues to search the public
5 databases showed some similarity to outer surface and membrane spanning proteins. Since the ORF3-18 product complemented the leaderless *phoA* gene, it can be concluded that this protein (or part thereof) would most probably be located extracellularly.

10 Example 11

A further clone was selected containing a plasmid designated *pho3-1*. This plasmid contained a gene (or part thereof), which complemented the leaderless *phoA*. The nucleotide and deduced amino acid sequences of the gene are
15 shown as SEQ ID NOS. 22 and 23.

A comparison of the amino acid sequence of *pho3-1* was performed.

Homologues to the GBS *pho3-1* gene product can be identified in *Streptococcus pyogenes*, *Streptococcus pneumoniae*, *Bacillus subtilis* (*yutD*) and *Enterococcus faecalis*. The *S. pyogenes*, *S. pneumoniae* and *E. faecalis*
20 homologues were identified from genome sequence data and no annotations were available as to the identity of the gene or gene products. In *B. subtilis*, the function of the *yutD* gene product is unknown. It can be noted however, that the *yutD* gene is located on the *B. subtilis* chromosome in a region containing genes involved in cell wall synthesis. The fact that this DNA sequence complemented the leaderless
25 *phoA* gene suggests that this gene product is extracellularly located.
30

Example 12

A further clone was selected containing a plasmid designated *pho3-21*. This plasmid contained a gene (or part thereof), which complemented the leaderless *phoA*. The
35 nucleotide and deduced amino acid sequences of the gene are shown as SEQ ID NOS. 24 and 25.

A comparison of the amino acid sequence of pho3-21 was performed.

Homologues to the GBS pho3-21 gene product can be identified in *Streptococcus pyogenes*, *Streptococcus pneumoniae*, *Lactobacillus fermentum* (bspA) and *Lactobacillus reuteri* (cnb). The *S. pyogenes* and *S. pneumoniae* homologues were identified from genome sequence data and no annotations were available as to the identity of the gene or gene products. In *L. fermentum*, the bspA gene product has been identified as being a basic cell surface-located protein that has some sequence similarity to family III of the bacterial solute-binding proteins (Turner, M.S. et al. 1997. J. Bacteriol. 179:3310-3316). In *L. reuteri*, the cnb gene product has been identified as a collagen binding protein that has some sequence similarity to the solute-binding component of bacterial ABC transporters (Roos, S. et al. 1996. FEMS Microbiol. Lett. 144:33-38).

Example 13

A further clone was selected containing a plasmid designated pho3-22. This plasmid contained a gene (or part thereof), which complemented the leaderless phoA. The nucleotide and deduced amino acid sequences of the gene are shown as SEQ ID NOS. 26 and 27.

A comparison of the amino acid sequence of pho3-22 was performed.

Homologues to the GBS pho3-22 gene product can be identified in *Enterococcus faecalis*, *Streptococcus equisimilis* (lppC), *Pseudomonas fluorescens* (oprI) and *Streptococcus thermophilus* (orf142). The *E. faecalis* homolog was identified from genome sequence data and no annotations were available as to the identity of the gene or gene products. In *S. equisimilis*, the lppC gene product has been identified as being a lipoprotein that is homologous to the E(P4) outer membrane protein from *Haemophilus influenzae* (Gase, K. et al. 1997. Med. Microbiol. Immunol. 186:63-73). Likewise, the *P.*

fluorescens *oprI* gene encodes a major outer membrane lipoprotein (Cornelis, P. et al. 1989. Mol. Microbiol. 3:421-428). In *S. thermophilus*, the *orf142* product has been putatively identified as a cell surface exposed lipoprotein that may act as a receptor for the bacteriophages TP-J34 and Sfi21 (Neve, H. et al. 1998. Virology 241:61-72). The ORF3-22 product showed good similarity to the above homologues, particularly at the N-terminus. This is most likely the region required for complementation of the leaderless *phoA* gene, and therefore serves as a leader sequence.

Example 14

A further clone was selected containing a plasmid designated *pho3-23*. This plasmid contained a gene (or part thereof), which complemented the leaderless *phoA*. The nucleotide and deduced amino acid sequences of the genes are shown as SEQ ID NOS. 28 and 29.

A comparison of the amino acid sequence of *pho3-23* was performed.

Homologues to the GBS *pho3-23* gene product can be identified in *Streptococcus pyogenes*, *Streptococcus pneumoniae*, *Enterococcus faecalis* and *Streptococcus mutans* (*perM*). The *S. pyogenes*, *S. pneumoniae* and *E. faecalis* homologues were identified from genome sequence data and no annotations were available as to the identity of the gene or gene products. In *S. mutans*, the *perM* gene product has been presumptively identified as a permease, but no other information is available as to the function of this protein. Considering that the *pho3-23* coding region complements the leaderless *phoA* gene, it can be concluded that the *pho3-17* gene product would most probably be located extracellularly.

Example 15

A further clone was selected containing a plasmid designated *pho3-24*. This plasmid contained a gene (or part thereof), which complemented the leaderless *phoA*. The

nucleotide and deduced amino acid sequences of the gene are shown as SEQ ID NOS. 30 and 31.

A comparison of the amino acid sequence of pho3-24 was performed.

5 Homologues to the GBS pho3-24 gene product can be identified in *Streptococcus mutans* (dltB), *Streptococcus pneumoniae*, *Streptococcus pyogenes*, *Enterococcus faecalis*, *Lactobacillus casei* (dltB) and *Bacillus subtilis* (dltB). The *S. pneumoniae*, *S. pyogenes* and *E. faecalis* homologues
10 were identified from genome sequence data and no annotations were available as to the identity of the gene or gene products. In *S. mutans*, *L. casei* and *B. subtilis*, the dltB gene product has been identified as being a basic membrane protein that is involved in the transport of
15 activated D-alanine through the cell membrane. The dltB gene product is involved in the biosynthesis of D-alanyl-lipoteichoic acid (Heaton, M.P. and F.C. Neuhaus. 1992. J. Bacteriol. 174:4707-4717). In *L. casei* and *B. subtilis*, the dltB gene product is believed to contain at least 9
20 membrane spanning domains, indicating that the protein or portions thereof are exposed to the outside of the cell.

Example 16

A further clone was selected containing a plasmid designated pho3-29. This plasmid contained a gene (or part
25 thereof), which complemented the leaderless phoA. The nucleotide and deduced amino acid sequences of the gene are shown as SEQ ID NOS. 32 and 33.

A comparison of the amino acid sequence of pho3-29 was performed.

30 Homologues to the GBS pho3-29 gene product can be identified in *Borrelia burgdorferi* (p23 or ospC), *Bacillus brevis* (owp) and *Pseudomonas aeruginosa* (oprI). Although these homologues are not related to each other, they all represent major outer surface proteins. In *B. burgdorferi*,
35 the ospC gene product has been identified as being a 23-kDa protein that is the immunodominant antigen on the surface of this bacterium (Padula, S.J. et al. 1993. Infect. Immun.

61:5097-5105). The *owp* gene product from *B. brevis* is one of two major cell wall proteins involved in the surface layer lattice (Tsuboi, A. 1988. J. Bacteriol. 170:935-945). Finally, the *oprI* gene from *P. aeruginosa* encodes a major outer membrane lipoprotein precursor (Saint-Onge, A. et al. 1992. J. Gen. Microbiol. 138:733-741).

Example 17

A further clone was selected containing a plasmid designated pho3-50. This plasmid contained a gene (or part thereof), which complemented the leaderless *phoA*. The nucleotide and deduced amino acid sequences of the gene are shown as SEQ ID NOS. 34 and 35.

A comparison of the amino acid sequence of pho3-50 was performed.

Homologues to the GBS pho3-50 gene product can be identified in a variety of Streptococci (*penA*, *pbp2B*, *pbpB2*), *Borrelia burgdorferi* (*pbp2*), *Enterococcus faecalis* (*pbpC*), *Staphylococcus aureus* (*pbpA*), *Mycobacterium leprae* (*pbpB*) and *Helicobacter pylori* (*pbp2*). In all cases, the above homologues can be identified as penicillin binding proteins (PBPs). Genes encoding penicillin binding proteins are often located in a cluster of genes associated with cell wall synthesis (Pucci, M.J. et al. 1997. J. Bacteriol. 179:5632-5635). Furthermore, PBPs are typically integrated into the cell wall of a bacterium with some or all of the protein being exposed on the outer bacterial surface.

CLAIMS

1. A peptide encoded by an operon including any of the genes identified herein as pho1-13, pho3-21, pho2-15, pho3-18, pho3-22, pho3-3, pho3-17, pho2-2, pho1-5, pho3-1, pho3-23, pho3-50, pho1-14, pho2-10, pho3-14, pho3-24 and pho3-29, obtainable from Group B *Streptococcus*, or a homologue thereof or a functional fragment thereof.
2. A peptide according to claim 1, comprising any of the amino acid sequences identified herein as SEQ ID NOS. 2, 4, 6, 8, 10, 11, 13, 15, 17, 19, 21, 23, 25, 27, 29, 31, 33 and 35.
3. A peptide according to claim 1 or claim 2, for therapeutic use.
4. A polynucleotide encoding a peptide according to claim 1 or claim 2, for therapeutic use.
5. A host transformed to express a peptide according to claim 1 or claim 2.
6. A vaccine comprising a peptide according to claim 1 or claim 2, or the means for its expression.
7. Use of a product according to any of claims 1 to 5, for screening potential drugs or for the detection of virulence.
8. Use of a product according to any of claims 1 to 5, for the manufacture of a medicament for use in the treatment or prevention of a condition associated with bacterial infection.
9. Use according to claim 8, wherein the infection is a Group B streptococcal infection.
10. Use according to claim 8 or claim 9, wherein the infection is a focal infection.
11. Use according to claim 8 or claim 9, wherein the infection is a urinary tract infection.
12. An antibody raised against a peptide according to claim 1 or claim 2.

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 Ile Asp Ala Ser Lys Gly Ile Phe Val Pro Arg Gly Val Ala Asn Gly
 115 120 125

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 145 150 155 160

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 Asp Pro Asn Leu Gly Ile Gln Trp Glu Asn Xaa Glu Glu Ala Glu Val
 165 170 175

tca gaa gca gat aag aat cac cca ctt ctc aaa gat gta aaa cct ttg 576
 Ser Glu Ala Asp Lys Asn His Pro Leu Leu Lys Asp Val Lys Pro Leu
 180 185 190

aag aag gaa gat ttg taa 594
 Lys Lys Glu Asp Leu
 195

<210> 8

<211> 197

<212> PRT

<213> group B streptococcus

<400> 8

Met Thr Glu Pro Phe Phe Asp Lys Glu Leu Thr Cys Arg Pro Ile Glu
 1 5 10 15

Ala Ile Pro Glu Leu Leu Glu Phe Asp Ile Thr Val Arg Gly Asp Asn
 20 25 30

Arg Gly Trp Phe Lys Glu Asn Phe Gln Lys Glu Lys Met Ile Pro Leu
 35 40 45

Gly Phe Pro Glu Ser Phe Phe Glu Ala Asp Lys Leu Gln Asn Asn Ile
 50 55 60

Ser Phe Thr Lys Lys Asn Thr Leu Arg Gly Leu His Ala Glu Pro Trp

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<400> 9
tat tat tta atc gga ggg ttg gca gaa atg caa cat gtc aat cat tct    48
Tyr Tyr Leu Ile Gly Gly Leu Ala Glu Met Gln His Val Asn His Ser
  1             5             10             15

tct ttt gat aaa gca tca aaa gca gga ttt att att gct tta ggc att    96
Ser Phe Asp Lys Ala Ser Lys Ala Gly Phe Ile Ile Ala Leu Gly Ile

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20	25	30	
gtt tat gga gat att ggt aca agc cca ctc tat acg atg caa tca ttg			144
Val Tyr Gly Asp Ile Gly Thr Ser Pro Leu Tyr Thr Met Gln Ser Leu			
35	40	45	
gtt gaa aac caa ggt ggt att tct agt gtc aca gaa tcg ttt atc tta			192
Val Glu Asn Gln Gly Gly Ile Ser Ser Val Thr Glu Ser Phe Ile Leu			
50	55	60	
ggg tct ata tct tta atc ata tgg acc ttg aca ctt att aca act atc			240
Gly Ser Ile Ser Leu Ile Ile Trp Thr Leu Thr Leu Ile Thr Thr Ile			
65	70	75	80
aag tat gtg ctt gta gct tta aag gcg gat aat cac cac gaa ggt ggt			288
Lys Tyr Val Leu Val Ala Leu Lys Ala Asp Asn His His Glu Gly Gly			
85	90	95	
att ttt tct tta tat acc ctt gtt aga aaa atg aca cct tgg tta att			336
Ile Phe Ser Leu Tyr Thr Leu Val Arg Lys Met Thr Pro Trp Leu Ile			
100	105	110	
gtt ccg gct gtt att gga ggt gca acc ttg ttg tca gat gga gct ttg			384
Val Pro Ala Val Ile Gly Gly Ala Thr Leu Leu Ser Asp Gly Ala Leu			
115	120	125	
acg cca gct gta acc gta ctt cag ccg tta agg att aaa gta gtt cct			432
Thr Pro Ala Val Thr Val Leu Gln Pro Leu Arg Ile Lys Val Val Pro			
130	135	140	
agt ttg cag cat att tcc aga atc aga gta tgt tat ttt gcg acc ttg			480
Ser Leu Gln His Ile Ser Arg Ile Arg Val Cys Tyr Phe Ala Thr Leu			
145	150	155	160
tta ttt act gtt act ttt gcc atc caa ggt ttg gaa cgg gtg tta ttg			528
Leu Phe Thr Val Thr Phe Ala Ile Gln Gly Leu Glu Arg Val Leu Leu			
165	170	175	
gaa tta ttg gcc att atg tta tat ggt ttg cct ttt ggt tta			570
Glu Leu Leu Ala Ile Met Leu Tyr Gly Leu Pro Phe Gly Leu			
180	185	190	
ncgggtctcct tatagttttg cccatccaga agttttcaag cattaatcca tactacggtt			630
tgaaattggtt atttagtcca gagaatcata aaggtatttt tatttttag gat cta ttt			687
		Asp Leu Phe	
tcc tgg cga caa acg gga gca gaa gca cta tac tct gac tta ggt cat			735

Ser Trp Arg Gln Thr Gly Ala Glu Ala Leu Tyr Ser Asp Leu Gly His
 195 200 205

ggt ggg cgt gga aat ata cat gtt tca tgg ccg ttc gtt aag gtt gcc 783
 Val Gly Arg Gly Asn Ile His Val Ser Trp Pro Phe Val Lys Val Ala
 210 215 220 225

att ata ctt tct tat tgt ggg caa ggg gca tgg att tta gct aat aag 831
 Ile Ile Leu Ser Tyr Cys Gly Gln Gly Ala Trp Ile Leu Ala Asn Lys
 230 235 240

aac gca gga aat gaa ttg aat ccc ttt ttt gct agt att cct tcg caa 879
 Asn Ala Gly Asn Glu Leu Asn Pro Phe Phe Ala Ser Ile Pro Ser Gln
 245 250 255

ttt aca atg cat gtc gtt att tta gct act ttg gca gct atc atc gct 927
 Phe Thr Met His Val Val Ile Leu Ala Thr Leu Ala Ala Ile Ile Ala
 260 265 270

tca cag gca ctg att tct ggatcaattt accttaagtt ctgagctatg 975
 Ser Gln Ala Leu Ile Ser
 275

cgactaaaaa tattcccaca atttcgttca acttatcctg ttgacaatat tgggtcaaac 1035

ctacatacct ggtattaatt ggttcttatt tgccattaca acctctattg gtttgctttt 1095

taagacttca gcgcacatgg aagcagcata tggattagcg ataacaatta cgatgcta 1155

gacaactatt ttactgtctt tctttttaat tcaaaaagga gtaaagagag gtttttagcta 1215

tt 1217

<210> 10

<211> 190

<212> PRT

<213> group B streptococcus

<400> 10

Tyr Tyr Leu Ile Gly Gly Leu Ala Glu Met Gln His Val Asn His Ser
 1 5 10 15

Ser Phe Asp Lys Ala Ser Lys Ala Gly Phe Ile Ile Ala Leu Gly Ile
 20 25 30

Val Tyr Gly Asp Ile Gly Thr Ser Pro Leu Tyr Thr Met Gln Ser Leu
 35 40 45

Val Glu Asn Gln Gly Gly Ile Ser Ser Val Thr Glu Ser Phe Ile Leu
 50 55 60
 Gly Ser Ile Ser Leu Ile Ile Trp Thr Leu Thr Leu Ile Thr Thr Ile
 65 70 75 80
 Lys Tyr Val Leu Val Ala Leu Lys Ala Asp Asn His His Glu Gly Gly
 85 90 95
 Ile Phe Ser Leu Tyr Thr Leu Val Arg Lys Met Thr Pro Trp Leu Ile
 100 105 110
 Val Pro Ala Val Ile Gly Gly Ala Thr Leu Leu Ser Asp Gly Ala Leu
 115 120 125
 Thr Pro Ala Val Thr Val Leu Gln Pro Leu Arg Ile Lys Val Val Pro
 130 135 140
 Ser Leu Gln His Ile Ser Arg Ile Arg Val Cys Tyr Phe Ala Thr Leu
 145 150 155 160
 Leu Phe Thr Val Thr Phe Ala Ile Gln Gly Leu Glu Arg Val Leu Leu
 165 170 175
 Glu Leu Leu Ala Ile Met Leu Tyr Gly Leu Pro Phe Gly Leu
 180 185 190

<210> 11

<211> 89

<212> PRT

<213> group B streptococcus

<400> 11

Asp Leu Phe Ser Trp Arg Gln Thr Gly Ala Glu Ala Leu Tyr Ser Asp
 1 5 10 15
 Leu Gly His Val Gly Arg Gly Asn Ile His Val Ser Trp Pro Phe Val
 20 25 30
 Lys Val Ala Ile Ile Leu Ser Tyr Cys Gly Gln Gly Ala Trp Ile Leu
 35 40 45
 Ala Asn Lys Asn Ala Gly Asn Glu Leu Asn Pro Phe Phe Ala Ser Ile
 50 55 60
 Pro Ser Gln Phe Thr Met His Val Val Ile Leu Ala Thr Leu Ala Ala

65

70

75

80

Ile Ile Ala Ser Gln Ala Leu Ile Ser

85

<210> 12

<211> 378

<212> DNA

<213> group B streptococcus

<220>

<221> CDS

<222> (1)..(378)

<400> 12

atg	cag	gta	ttt	tta	aat	att	gtc	aat	aaa	ttc	ttt	gat	cca	ggt	att	48
Met	Gln	Val	Phe	Leu	Asn	Ile	Val	Asn	Lys	Phe	Phe	Asp	Pro	Val	Ile	
1				5					10					15		

cat	atg	ggt	tcg	gga	ggt	gtg	atg	cta	att	gtc	atg	aca	ggt	tta	gcc	96
His	Met	Gly	Ser	Gly	Val	Val	Met	Leu	Ile	Val	Met	Thr	Gly	Leu	Ala	
			20					25					30			

atg	ata	ttt	gga	gtg	aag	ttt	tct	aaa	gca	ctt	gaa	ggt	ggt	att	aag	144
Met	Ile	Phe	Gly	Val	Lys	Phe	Ser	Lys	Ala	Leu	Glu	Gly	Gly	Ile	Lys	
		35					40					45				

tta	gct	att	gct	ctt	acg	ggt	att	ggt	gct	att	att	ggt	att	tta	act	192
Leu	Ala	Ile	Ala	Leu	Thr	Gly	Ile	Gly	Ala	Ile	Ile	Gly	Ile	Leu	Thr	
	50					55					60					

ggt	gct	ttt	tcc	gaa	tca	ctt	caa	gct	ttt	ggt	aaa	aat	aca	gga	atc	240
Gly	Ala	Phe	Ser	Glu	Ser	Leu	Gln	Ala	Phe	Val	Lys	Asn	Thr	Gly	Ile	
65					70					75				80		

aat	cta	agc	att	att	gac	ggt	ggt	tgg	gct	cca	tta	gca	act	att	aca	288
Asn	Leu	Ser	Ile	Ile	Asp	Val	Gly	Trp	Ala	Pro	Leu	Ala	Thr	Ile	Thr	
			85					90					95			

tgg	gga	tca	cca	tat	acg	ctt	tac	ttc	tta	tta	atc	atg	ctt	att	gtc	336
Trp	Gly	Ser	Pro	Tyr	Thr	Leu	Tyr	Phe	Leu	Leu	Ile	Met	Leu	Ile	Val	
			100					105					110			

aat	att	ggt	atg	att	ggt	atg	aaa	aaa	aaa	cgg	ata	cct	tag			378
Asn	Ile	Val	Met	Ile	Val	Met	Lys	Lys	Lys	Arg	Ile	Pro				
		115					120					125				

<210> 13

<211> 125

<212> PRT

<213> group B streptococcus

<400> 13

Met Gln Val Phe Leu Asn Ile Val Asn Lys Phe Phe Asp Pro Val Ile
 1 5 10 15

His Met Gly Ser Gly Val Val Met Leu Ile Val Met Thr Gly Leu Ala
 20 25 30

Met Ile Phe Gly Val Lys Phe Ser Lys Ala Leu Glu Gly Gly Ile Lys
 35 40 45

Leu Ala Ile Ala Leu Thr Gly Ile Gly Ala Ile Ile Gly Ile Leu Thr
 50 55 60

Gly Ala Phe Ser Glu Ser Leu Gln Ala Phe Val Lys Asn Thr Gly Ile
 65 70 75 80

Asn Leu Ser Ile Ile Asp Val Gly Trp Ala Pro Leu Ala Thr Ile Thr
 85 90 95

Trp Gly Ser Pro Tyr Thr Leu Tyr Phe Leu Leu Ile Met Leu Ile Val
 100 105 110

Asn Ile Val Met Ile Val Met Lys Lys Lys Arg Ile Pro
 115 120 125

<210> 14

<211> 705

<212> DNA

<213> group B streptococcus

<220>

<221> CDS

<222> (118)..(705)

<400> 14

ggatcgggcg caagcttaac gattctttttt aaaatcatta aattttaaaa caaatttcag 60

acatattgcc aaagttttga tattattact ataatatagt ttgtagagga gaataat 117

atg ggc caa gaa cct atc atc gaa tat caa aat atc aat aaa gtg tat	165
Met Gly Gln Glu Pro Ile Ile Glu Tyr Gln Asn Ile Asn Lys Val Tyr	
1 5 10 15	
ggg gaa aat gtt gcg gtt gaa gat att aac ctt aaa att tac cct ggt	213
Gly Glu Asn Val Ala Val Glu Asp Ile Asn Leu Lys Ile Tyr Pro Gly	
20 25 30	
gat ttc gtt tgt ttc atc ggt acg agt gga tca ggt aaa aca aca tta	261
Asp Phe Val Cys Phe Ile Gly Thr Ser Gly Ser Gly Lys Thr Thr Leu	
35 40 45	
atg cgt atg gtt aac cat atg tta aaa cca aca aat ggt act cta tta	309
Met Arg Met Val Asn His Met Leu Lys Pro Thr Asn Gly Thr Leu Leu	
50 55 60	
ttt aag gga aaa gat atc tct act att aac ccc att gaa tta aga cgc	357
Phe Lys Gly Lys Asp Ile Ser Thr Ile Asn Pro Ile Glu Leu Arg Arg	
65 70 75 80	
aga att gga tat gtt atc caa aac att ggt tta atg cct cat atg acc	405
Arg Ile Gly Tyr Val Ile Gln Asn Ile Gly Leu Met Pro His Met Thr	
85 90 95	
att tac gaa aat ata gtt ctt gta cca aaa tta ttg aaa tgg tca gaa	453
Ile Tyr Glu Asn Ile Val Leu Val Pro Lys Leu Leu Lys Trp Ser Glu	
100 105 110	
gaa gct aaa aga gct aaa gca agg gaa ctt att aaa tta gtt gaa tta	501
Glu Ala Lys Arg Ala Lys Ala Arg Glu Leu Ile Lys Leu Val Glu Leu	
115 120 125	
ccc gaa gaa tat ttg gat cgc tac cct agt gag ttg tct ggc ggt cag	549
Pro Glu Glu Tyr Leu Asp Arg Tyr Pro Ser Glu Leu Ser Gly Gly Gln	
130 135 140	
caa caa cgt atc ggt gtc att cgc gct ctt gca gca gac caa gat att	597
Gln Gln Arg Ile Gly Val Ile Arg Ala Leu Ala Ala Asp Gln Asp Ile	
145 150 155 160	
att tta atg gat gag cct ttt gga gct ctg gat cct att act aga gaa	645
Ile Leu Met Asp Glu Pro Phe Gly Ala Leu Asp Pro Ile Thr Arg Glu	
165 170 175	
ggt att caa gac ttt agt caa gtc tct tca gga aga aat ggg gga aaa	693
Gly Ile Gln Asp Phe Ser Gln Val Ser Ser Gly Arg Asn Gly Gly Lys	
180 185 190	

cta tca tct tag
 Leu Ser Ser
 195

705

<210> 15
 <211> 195
 <212> PRT
 <213> group B streptococcus

<400> 15
 Met Gly Gln Glu Pro Ile Ile Glu Tyr Gln Asn Ile Asn Lys Val Tyr
 1 5 10 15

Gly Glu Asn Val Ala Val Glu Asp Ile Asn Leu Lys Ile Tyr Pro Gly
 20 25 30

Asp Phe Val Cys Phe Ile Gly Thr Ser Gly Ser Gly Lys Thr Thr Leu
 35 40 45

Met Arg Met Val Asn His Met Leu Lys Pro Thr Asn Gly Thr Leu Leu
 50 55 60

Phe Lys Gly Lys Asp Ile Ser Thr Ile Asn Pro Ile Glu Leu Arg Arg
 65 70 75 80

Arg Ile Gly Tyr Val Ile Gln Asn Ile Gly Leu Met Pro His Met Thr
 85 90 95

Ile Tyr Glu Asn Ile Val Leu Val Pro Lys Leu Leu Lys Trp Ser Glu
 100 105 110

Glu Ala Lys Arg Ala Lys Ala Arg Glu Leu Ile Lys Leu Val Glu Leu
 115 120 125

Pro Glu Glu Tyr Leu Asp Arg Tyr Pro Ser Glu Leu Ser Gly Gly Gln
 130 135 140

Gln Gln Arg Ile Gly Val Ile Arg Ala Leu Ala Ala Asp Gln Asp Ile
 145 150 155 160

Ile Leu Met Asp Glu Pro Phe Gly Ala Leu Asp Pro Ile Thr Arg Glu
 165 170 175

Gly Ile Gln Asp Phe Ser Gln Val Ser Ser Gly Arg Asn Gly Gly Lys
 180 185 190

Leu Ser Ser

195

<210> 16

<211> 367

<212> DNA

<213> group B streptococcus

<220>

<221> CDS

<222> (1)..(366)

<400> 16

atc cct tat agt gat gtt ttt gct aca gga gga ttt tta tac tat gta	48
Ile Pro Tyr Ser Asp Val Phe Ala Thr Gly Gly Phe Leu Tyr Tyr Val	
1 5 10 15	

acg att gct cta agt tac ctt tta ggg tct agt atc tgg tta ttt att	96
Thr Ile Ala Leu Ser Tyr Leu Leu Gly Ser Ser Ile Trp Leu Phe Ile	
20 25 30	

gta cag ttt att gct tac tat gta tct gga att tat ttt tat aaa tta	144
Val Gln Phe Ile Ala Tyr Tyr Val Ser Gly Ile Tyr Phe Tyr Lys Leu	
35 40 45	

gtt tat tat gtg gca caa agt gaa att gtc tcg ata ggc atg acg ttg	192
Val Tyr Tyr Val Ala Gln Ser Glu Ile Val Ser Ile Gly Met Thr Leu	
50 55 60	

att ttc tat ata atg aat att gtc tta gga ttc ggt ggt atg tac cca	240
Ile Phe Tyr Ile Met Asn Ile Val Leu Gly Phe Gly Gly Met Tyr Pro	
65 70 75 80	

ata cag tgg gca tta cct ttt atg ctc att tcg cta tgg ttt tta att	288
Ile Gln Trp Ala Leu Pro Phe Met Leu Ile Ser Leu Trp Phe Leu Ile	
85 90 95	

aaa ttt tgt gtc gat aat atc gtt gat gaa gca ttt ata ttt tat ggt	336
Lys Phe Cys Val Asp Asn Ile Val Asp Glu Ala Phe Ile Phe Tyr Gly	
100 105 110	

att tta gca gca ttc tca cta ttt ata gat c	367
Ile Leu Ala Ala Phe Ser Leu Phe Ile Asp	
115 120	

<210> 17

<211> 122

<212> PRT

<213> group B streptococcus

<400> 17

Ile Pro Tyr Ser Asp Val Phe Ala Thr Gly Gly Phe Leu Tyr Tyr Val
 1 5 10 15

Thr Ile Ala Leu Ser Tyr Leu Leu Gly Ser Ser Ile Trp Leu Phe Ile
 20 25 30

Val Gln Phe Ile Ala Tyr Tyr Val Ser Gly Ile Tyr Phe Tyr Lys Leu
 35 40 45

Val Tyr Tyr Val Ala Gln Ser Glu Ile Val Ser Ile Gly Met Thr Leu
 50 55 60

Ile Phe Tyr Ile Met Asn Ile Val Leu Gly Phe Gly Gly Met Tyr Pro
 65 70 75 80

Ile Gln Trp Ala Leu Pro Phe Met Leu Ile Ser Leu Trp Phe Leu Ile
 85 90 95

Lys Phe Cys Val Asp Asn Ile Val Asp Glu Ala Phe Ile Phe Tyr Gly
 100 105 110

Ile Leu Ala Ala Phe Ser Leu Phe Ile Asp
 115 120

<210> 18

<211> 570

<212> DNA

<213> group B streptococcus

<220>

<221> CDS

<222> (1)..(570)

<400> 18

atg agg aaa cgt ttt tcc ttg cta aat ttt att gtt gtt act ttt att 48
 Met Arg Lys Arg Phe Ser Leu Leu Asn Phe Ile Val Val Thr Phe Ile
 1 5 10 15

ttc ttt ttc ttt att ctt ttt ccg ctt tta aac cat aag gga aaa gta 96
 Phe Phe Phe Phe Ile Leu Phe Pro Leu Leu Asn His Lys Gly Lys Val
 20 25 30

gat gct aat tct agg cag agt gtt acc tac acc aaa gaa gaa ttt ata 144
 Asp Ala Asn Ser Arg Gln Ser Val Thr Tyr Thr Lys Glu Glu Phe Ile
 35 40 45

caa aaa att gtg cca gat gcg caa gat cta gga aag tcg tac ggt att 192
 Gln Lys Ile Val Pro Asp Ala Gln Asp Leu Gly Lys Ser Tyr Gly Ile
 50 55 60

cgt cct tca ttt att att gca cag gcg gct ttg gat tct gat ttc gga 240
 Arg Pro Ser Phe Ile Ile Ala Gln Ala Ala Leu Asp Ser Asp Phe Gly
 65 70 75 80

gag aaa tat agc tat agt atc ata atc tgt tgg ttg ctt gca gaa cca 288
 Glu Lys Tyr Ser Tyr Ser Ile Ile Ile Cys Trp Leu Leu Ala Glu Pro
 85 90 95

gga acg ccc tca att acc tta aat gat agt agt aca gga aaa aaa cag 336
 Gly Thr Pro Ser Ile Thr Leu Asn Asp Ser Ser Thr Gly Lys Lys Gln
 100 105 110

gaa aag caa ttt act cat tat aaa tct tgg aag tat tca atg gat gat 384
 Glu Lys Gln Phe Thr His Tyr Lys Ser Trp Lys Tyr Ser Met Asp Asp
 115 120 125

tac ctt gct cat ata aaa tct gga gcg aca ggc aaa aaa gat tca tat 432
 Tyr Leu Ala His Ile Lys Ser Gly Ala Thr Gly Lys Lys Asp Ser Tyr
 130 135 140

act ata atg gtg tct gtt aaa aat cca aaa act tta gtg caa aaa tta 480
 Thr Ile Met Val Ser Val Lys Asn Pro Lys Thr Leu Val Gln Lys Leu
 145 150 155 160

caa gat agt ggt ttt gat aat gac aaa aag tac gct aaa aaa atg acg 528
 Gln Asp Ser Gly Phe Asp Asn Asp Lys Lys Tyr Ala Lys Lys Met Thr
 165 170 175

gaa atc att gat ttg tat gat tta aca aga tat gat aag tga 570
 Glu Ile Ile Asp Leu Tyr Asp Leu Thr Arg Tyr Asp Lys
 180 185 190

<210> 19

<211> 189

<212> PRT

<213> group B streptococcus

<400> 19

Met Arg Lys Arg Phe Ser Leu Leu Asn Phe Ile Val Val Thr Phe Ile
 1 5 10 15
 Phe Phe Phe Phe Ile Leu Phe Pro Leu Leu Asn His Lys Gly Lys Val
 20 25 30
 Asp Ala Asn Ser Arg Gln Ser Val Thr Tyr Thr Lys Glu Glu Phe Ile
 35 40 45
 Gln Lys Ile Val Pro Asp Ala Gln Asp Leu Gly Lys Ser Tyr Gly Ile
 50 55 60
 Arg Pro Ser Phe Ile Ile Ala Gln Ala Ala Leu Asp Ser Asp Phe Gly
 65 70 75 80
 Glu Lys Tyr Ser Tyr Ser Ile Ile Ile Cys Trp Leu Leu Ala Glu Pro
 85 90 95
 Gly Thr Pro Ser Ile Thr Leu Asn Asp Ser Ser Thr Gly Lys Lys Gln
 100 105 110
 Glu Lys Gln Phe Thr His Tyr Lys Ser Trp Lys Tyr Ser Met Asp Asp
 115 120 125
 Tyr Leu Ala His Ile Lys Ser Gly Ala Thr Gly Lys Lys Asp Ser Tyr
 130 135 140
 Thr Ile Met Val Ser Val Lys Asn Pro Lys Thr Leu Val Gln Lys Leu
 145 150 155 160
 Gln Asp Ser Gly Phe Asp Asn Asp Lys Lys Tyr Ala Lys Lys Met Thr
 165 170 175
 Glu Ile Ile Asp Leu Tyr Asp Leu Thr Arg Tyr Asp Lys
 180 185

<210> 20

<211> 978

<212> DNA

<213> group B streptococcus

<220>

<221> CDS

<222> (1)..(978)

<400> 20

atg ctt gtc atc att ttg atc att gta cta gct agt ctg aca gtg acg	48
Met Leu Val Ile Ile Leu Ile Ile Val Leu Ala Ser Leu Thr Val Thr	
1 5 10 15	
ata att tct tac cca aaa atg acg gaa tta aca aag tcc gtt gaa aaa	96
Ile Ile Ser Tyr Pro Lys Met Thr Glu Leu Thr Lys Ser Val Glu Lys	
20 25 30	
caa ctt gaa gat aat gct gat aat cta tca gac caa ctg aca tat cag	144
Gln Leu Glu Asp Asn Ala Asp Asn Leu Ser Asp Gln Leu Thr Tyr Gln	
35 40 45	
ata gaa gtg gcg caa aaa gat caa atc tac gtg act aat cag cta aac	192
Ile Glu Val Ala Gln Lys Asp Gln Ile Tyr Val Thr Asn Gln Leu Asn	
50 55 60	
cgt atg caa cag gaa att atc agt cgc tta ccg ata tgc gta cag aat	240
Arg Met Gln Gln Glu Ile Ile Ser Arg Leu Pro Ile Cys Val Gln Asn	
65 70 75 80	
aaa tca gca tta acg gag agt cga gat cga tca gac aaa cgc ttg gaa	288
Lys Ser Ala Leu Thr Glu Ser Arg Asp Arg Ser Asp Lys Arg Leu Glu	
85 90 95	
ttg att aac tcc aat tta tct cag tca gtt cag aaa atg caa gat tca	336
Leu Ile Asn Ser Asn Leu Ser Gln Ser Val Gln Lys Met Gln Asp Ser	
100 105 110	
atg aaa aac gct tgg atc aaa tgc gcc aaa ctg ttg agg aaa agc tgg	384
Met Lys Asn Ala Trp Ile Lys Cys Ala Lys Leu Leu Arg Lys Ser Trp	
115 120 125	
aaa aaa cgc tac aaa cgc gtt gca aac ttc ttt gaa act gta tcg cgt	432
Lys Lys Arg Tyr Lys Arg Val Ala Asn Phe Phe Glu Thr Val Ser Arg	
130 135 140	
caa cta gag agc gtc aat caa ggt ctg ggt aga tgg aaa ctg tgc caa	480
Gln Leu Glu Ser Val Asn Gln Gly Leu Gly Arg Trp Lys Leu Cys Gln	
145 150 155 160	
gat gtt ggt acc act gaa caa agt ctg tca aat act aag aca agg gga	528
Asp Val Gly Thr Thr Glu Gln Ser Leu Ser Asn Thr Lys Thr Arg Gly	
165 170 175	
ata tta ggg gag tta caa ctc ggt caa att ata gaa gat att atg aca	576
Ile Leu Gly Glu Leu Gln Leu Gly Gln Ile Ile Glu Asp Ile Met Thr	
180 185 190	

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gtt agt caa tat gag aga gaa ttt cct acg gtg tct ggc tct tct gag   624
Val Ser Gln Tyr Glu Arg Glu Phe Pro Thr Val Ser Gly Ser Ser Glu
      195                200                205

cgt gtt gaa tat gct att aaa tac ctg gaa atg gtc agg gag att ata   672
Arg Val Glu Tyr Ala Ile Lys Tyr Leu Glu Met Val Arg Glu Ile Ile
      210                215                220

tct att tgc cta ttg act cta agt ttc tct aga aga tta tta ccg att   720
Ser Ile Cys Leu Leu Thr Leu Ser Phe Ser Arg Arg Leu Leu Pro Ile
      225                230                235                240

ggg aga tgc tta tgg aat tgg gtg acc agg ttc aaa tgg aac tct att   768
Gly Arg Cys Leu Trp Asn Trp Val Thr Arg Phe Lys Trp Asn Ser Ile
      245                250                255

cgt aat ctt tac tgg gca agt att cgt aaa ttt gca aaa gat ata aac   816
Arg Asn Leu Tyr Trp Ala Ser Ile Arg Lys Phe Ala Lys Asp Ile Asn
      260                265                270

aat aag tac tta aat cct cct gaa acg aca aat ttt ggt atc atg ttc   864
Asn Lys Tyr Leu Asn Pro Pro Glu Thr Thr Asn Phe Gly Ile Met Phe
      275                280                285

tta cca act gaa ggg ctc tat tct gaa gtg gta aga aat gca aca ttc   912
Leu Pro Thr Glu Gly Leu Tyr Ser Glu Val Val Arg Asn Ala Thr Phe
      290                295                300

ttt gat agt cta aga cgt gac gaa aat att gta gta gct gga ccg tca   960
Phe Asp Ser Leu Arg Arg Asp Glu Asn Ile Val Val Ala Gly Pro Ser
      305                310                315                320

acc tta tct gct tac taa                                           978
Thr Leu Ser Ala Tyr
      325

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<210> 21

<211> 325

<212> PRT

<213> group B streptococcus

<400> 21

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Met Leu Val Ile Ile Leu Ile Ile Val Leu Ala Ser Leu Thr Val Thr
  1                5                10                15

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Ile Ile Ser Tyr Pro Lys Met Thr Glu Leu Thr Lys Ser Val Glu Lys
      20                25                30

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Gln Leu Glu Asp Asn Ala Asp Asn Leu Ser Asp Gln Leu Thr Tyr Gln
 35 40 45
 Ile Glu Val Ala Gln Lys Asp Gln Ile Tyr Val Thr Asn Gln Leu Asn
 50 55 60
 Arg Met Gln Gln Glu Ile Ile Ser Arg Leu Pro Ile Cys Val Gln Asn
 65 70 75 80
 Lys Ser Ala Leu Thr Glu Ser Arg Asp Arg Ser Asp Lys Arg Leu Glu
 85 90 95
 Leu Ile Asn Ser Asn Leu Ser Gln Ser Val Gln Lys Met Gln Asp Ser
 100 105 110
 Met Lys Asn Ala Trp Ile Lys Cys Ala Lys Leu Leu Arg Lys Ser Trp
 115 120 125
 Lys Lys Arg Tyr Lys Arg Val Ala Asn Phe Phe Glu Thr Val Ser Arg
 130 135 140
 Gln Leu Glu Ser Val Asn Gln Gly Leu Gly Arg Trp Lys Leu Cys Gln
 145 150 155 160
 Asp Val Gly Thr Thr Glu Gln Ser Leu Ser Asn Thr Lys Thr Arg Gly
 165 170 175
 Ile Leu Gly Glu Leu Gln Leu Gly Gln Ile Ile Glu Asp Ile Met Thr
 180 185 190
 Val Ser Gln Tyr Glu Arg Glu Phe Pro Thr Val Ser Gly Ser Ser Glu
 195 200 205
 Arg Val Glu Tyr Ala Ile Lys Tyr Leu Glu Met Val Arg Glu Ile Ile
 210 215 220
 Ser Ile Cys Leu Leu Thr Leu Ser Phe Ser Arg Arg Leu Leu Pro Ile
 225 230 235 240
 Gly Arg Cys Leu Trp Asn Trp Val Thr Arg Phe Lys Trp Asn Ser Ile
 245 250 255
 Arg Asn Leu Tyr Trp Ala Ser Ile Arg Lys Phe Ala Lys Asp Ile Asn
 260 265 270
 Asn Lys Tyr Leu Asn Pro Pro Glu Thr Thr Asn Phe Gly Ile Met Phe
 275 280 285

Leu Pro Thr Glu Gly Leu Tyr Ser Glu Val Val Arg Asn Ala Thr Phe
 290 295 300

Phe Asp Ser Leu Arg Arg Asp Glu Asn Ile Val Val Ala Gly Pro Ser
 305 310 315 320

Thr Leu Ser Ala Tyr
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<210> 22

<211> 579

<212> DNA

<213> group B streptococcus

<220>

<221> CDS

<222> (1)..(579)

<400> 22

atg cga aaa gaa gtg aca cca gag atg ctt aac tat aat aag tat cct 48
 Met Arg Lys Glu Val Thr Pro Glu Met Leu Asn Tyr Asn Lys Tyr Pro
 1 5 10 15

ggc cca cag ttt att cac ttt gaa aat atc gtt aaa agt gat gat att 96
 Gly Pro Gln Phe Ile His Phe Glu Asn Ile Val Lys Ser Asp Asp Ile
 20 25 30

gaa ttt caa ctt gtt att aat gaa aaa tca gct ttt gat gtg act gtc 144
 Glu Phe Gln Leu Val Ile Asn Glu Lys Ser Ala Phe Asp Val Thr Val
 35 40 45

ttt gga caa cgt ttt tct gag att tta tta aaa tat gat ttt atc gtt 192
 Phe Gly Gln Arg Phe Ser Glu Ile Leu Leu Lys Tyr Asp Phe Ile Val
 50 55 60

ggc gat tgg ggt aac gag cag ttg agg cta aga ggc ttt tac aaa gat 240
 Gly Asp Trp Gly Asn Glu Gln Leu Arg Leu Arg Gly Phe Tyr Lys Asp
 65 70 75 80

gct agt acg att aga aaa aat agc cgg att tca cgt tta gaa gat tat 288
 Ala Ser Thr Ile Arg Lys Asn Ser Arg Ile Ser Arg Leu Glu Asp Tyr
 85 90 95

att aaa gag tat tgt aac ttt ggt tgt gct tat ttt gtg ttg gag aat 336
 Ile Lys Glu Tyr Cys Asn Phe Gly Cys Ala Tyr Phe Val Leu Glu Asn

100	105	110	
cca aat cct aga gat att aaa ttt gat gat gaa aga cct cat aag cgt			384
Pro Asn Pro Arg Asp Ile Lys Phe Asp Asp Glu Arg Pro His Lys Arg			
115	120	125	
cgt aag tca aga tcc aaa tca caa tca tca aag tca caa act aga aat			432
Arg Lys Ser Arg Ser Lys Ser Gln Ser Ser Lys Ser Gln Thr Arg Asn			
130	135	140	
aat cgt tcc cag tca aat gcc aat gct cat ttt aca agt aaa aag cgt			480
Asn Arg Ser Gln Ser Asn Ala Asn Ala His Phe Thr Ser Lys Lys Arg			
145	150	155	160
aaa gac aca aaa cgc cgt caa gaa cgt cat att aaa gaa gag caa gat			528
Lys Asp Thr Lys Arg Arg Gln Glu Arg His Ile Lys Glu Glu Gln Asp			
165	170	175	
aag gaa atg acc tct gca aag cag cat ttg tta ttc gta aga aaa aat			576
Lys Glu Met Thr Ser Ala Lys Gln His Leu Leu Phe Val Arg Lys Asn			
180	185	190	
taa			579

<210> 23

<211> 192

<212> PRT

<213> group B streptococcus

<400> 23

Met Arg Lys Glu Val Thr Pro Glu Met Leu Asn Tyr Asn Lys Tyr Pro
1 5 10 15

Gly Pro Gln Phe Ile His Phe Glu Asn Ile Val Lys Ser Asp Asp Ile
20 25 30

Glu Phe Gln Leu Val Ile Asn Glu Lys Ser Ala Phe Asp Val Thr Val
35 40 45

Phe Gly Gln Arg Phe Ser Glu Ile Leu Leu Lys Tyr Asp Phe Ile Val
50 55 60

Gly Asp Trp Gly Asn Glu Gln Leu Arg Leu Arg Gly Phe Tyr Lys Asp
65 70 75 80

Ala Ser Thr Ile Arg Lys Asn Ser Arg Ile Ser Arg Leu Glu Asp Tyr

	85		90		95
Ile Lys Glu Tyr Cys Asn Phe Gly Cys Ala Tyr Phe Val Leu Glu Asn					
	100		105		110
Pro Asn Pro Arg Asp Ile Lys Phe Asp Asp Glu Arg Pro His Lys Arg					
	115		120		125
Arg Lys Ser Arg Ser Lys Ser Gln Ser Ser Lys Ser Gln Thr Arg Asn					
	130		135		140
Asn Arg Ser Gln Ser Asn Ala Asn Ala His Phe Thr Ser Lys Lys Arg					
	145		150		155
					160
Lys Asp Thr Lys Arg Arg Gln Glu Arg His Ile Lys Glu Glu Gln Asp					
	165		170		175
Lys Glu Met Thr Ser Ala Lys Gln His Leu Leu Phe Val Arg Lys Asn					
	180		185		190

<210> 24

<211> 609

<212> DNA

<213> group B streptococcus

<220>

<221> CDS

<222> (1)..(609)

<400> 24

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Met Thr Ile Lys Lys Val Leu Ser Val Thr Gly Ile Ile Leu Val Thr	
1 5 10 15	

gta gcg tct cta gct gct tgt agc tca aaa tct cat act act aag acg	96
Val Ala Ser Leu Ala Ala Cys Ser Ser Lys Ser His Thr Thr Lys Thr	
20 25 30	

ggc aaa aaa gaa gtt aat ttt gca act gtt gga aca acg gca cct ttt	144
Gly Lys Lys Glu Val Asn Phe Ala Thr Val Gly Thr Thr Ala Pro Phe	
35 40 45	

tct tat gtg aag gat ggg aaa ctg act ggc ttt gat att gaa gta gcc	192
Ser Tyr Val Lys Asp Gly Lys Leu Thr Gly Phe Asp Ile Glu Val Ala	
50 55 60	

aaa gct gtt ttt aaa ggt tca gat aac tat aaa gtc act ttt aaa aaa 240
 Lys Ala Val Phe Lys Gly Ser Asp Asn Tyr Lys Val Thr Phe Lys Lys
 65 70 75 80

aca gaa tgg tca tcg gta ttt acc ggc att gat tca gga aag ttt caa 288
 Thr Glu Trp Ser Ser Val Phe Thr Gly Ile Asp Ser Gly Lys Phe Gln
 85 90 95

atg ggt gga aat aat att tct tat tca tca gag aga tct caa aaa tay 336
 Met Gly Gly Asn Asn Ile Ser Tyr Ser Ser Glu Arg Ser Gln Lys Tyr
 100 105 110

tta ttt tca tac cca ata ggc tct act cct tca gtt tta gca gtt cct 384
 Leu Phe Ser Tyr Pro Ile Gly Ser Thr Pro Ser Val Leu Ala Val Pro
 115 120 125

aag aat agt aat atc aaa gct tat aat gat att agt ggt cat aaa aca 432
 Lys Asn Ser Asn Ile Lys Ala Tyr Asn Asp Ile Ser Gly His Lys Thr
 130 135 140

cag gtt gtc caa gga acg aca act gcc aag caa tta gaa aat ttc aat 480
 Gln Val Val Gln Gly Thr Thr Thr Ala Lys Gln Leu Glu Asn Phe Asn
 145 150 155 160

aaa gag cat cag aaa aat cct gtt act cta aaa tat act aat gaa aat 528
 Lys Glu His Gln Lys Asn Pro Val Thr Leu Lys Tyr Thr Asn Glu Asn
 165 170 175

att aca cag att cta acg aat ttg agt gat gga aaa gct gat ttt aaa 576
 Ile Thr Gln Ile Leu Thr Asn Leu Ser Asp Gly Lys Ala Asp Phe Lys
 180 185 190

ctt ttg acg gac caa ctg tta acg cta tta taa 609
 Leu Leu Thr Asp Gln Leu Leu Thr Leu Leu
 195 200

<210> 25

<211> 202

<212> PRT

<213> group B streptococcus

<400> 25

Met Thr Ile Lys Lys Val Leu Ser Val Thr Gly Ile Ile Leu Val Thr
 1 5 10 15

Val Ala Ser Leu Ala Ala Cys Ser Ser Lys Ser His Thr Thr Lys Thr
 20 25 30

Gly Lys Lys Glu Val Asn Phe Ala Thr Val Gly Thr Thr Ala Pro Phe
 35 40 45
 Ser Tyr Val Lys Asp Gly Lys Leu Thr Gly Phe Asp Ile Glu Val Ala
 50 55 60
 Lys Ala Val Phe Lys Gly Ser Asp Asn Tyr Lys Val Thr Phe Lys Lys
 65 70 75 80
 Thr Glu Trp Ser Ser Val Phe Thr Gly Ile Asp Ser Gly Lys Phe Gln
 85 90 95
 Met Gly Gly Asn Asn Ile Ser Tyr Ser Ser Glu Arg Ser Gln Lys Tyr
 100 105 110
 Leu Phe Ser Tyr Pro Ile Gly Ser Thr Pro Ser Val Leu Ala Val Pro
 115 120 125
 Lys Asn Ser Asn Ile Lys Ala Tyr Asn Asp Ile Ser Gly His Lys Thr
 130 135 140
 Gln Val Val Gln Gly Thr Thr Thr Ala Lys Gln Leu Glu Asn Phe Asn
 145 150 155 160
 Lys Glu His Gln Lys Asn Pro Val Thr Leu Lys Tyr Thr Asn Glu Asn
 165 170 175
 Ile Thr Gln Ile Leu Thr Asn Leu Ser Asp Gly Lys Ala Asp Phe Lys
 180 185 190
 Leu Leu Thr Asp Gln Leu Leu Thr Leu Leu
 195 200

<210> 26

<211> 357

<212> DNA

<213> group B streptococcus

<220>

<221> CDS

<222> (1)..(357)

<400> 26

atg aag aat ata aca aag cta tca act gtt gct tta agc cta cta ctt 48
 Met Lys Asn Ile Thr Lys Leu Ser Thr Val Ala Leu Ser Leu Leu Leu

1	5	10	15	
tgt acg gcg tgt gct gca tca aac acg tct aca tct aaa aca cag tct				96
Cys Thr Ala Cys Ala Ala Ser Asn Thr Ser Thr Ser Lys Thr Gln Ser				
20	25	30		
cat cat cct aaa caa act aaa ctc aca gat aag caa aaa gaa gaa ccc				144
His His Pro Lys Gln Thr Lys Leu Thr Asp Lys Gln Lys Glu Glu Pro				
35	40	45		
aaa aac aaa gaa gct gct gat caa gag atg cat ccc caa ggc gct gtt				192
Lys Asn Lys Glu Ala Ala Asp Gln Glu Met His Pro Gln Gly Ala Val				
50	55	60		
gat ttg aca aaa tat aag gca aaa ccg gtc aaa gat tat gga aaa aaa				240
Asp Leu Thr Lys Tyr Lys Ala Lys Pro Val Lys Asp Tyr Gly Lys Lys				
65	70	75	80	
atc gat gtt ggt gat ggc aag aaa atg aac att tat gaa act ggt cag				288
Ile Asp Val Gly Asp Gly Lys Lys Met Asn Ile Tyr Glu Thr Gly Gln				
85	90	95		
gga aaa att cca att gtt ttt att cct ggt caa gct gag att cgc cac				336
Gly Lys Ile Pro Ile Val Phe Ile Pro Gly Gln Ala Glu Ile Arg His				
100	105	110		
gct atg ctt ata aga att taa				357
Ala Met Leu Ile Arg Ile				
115				

<210> 27

<211> 118

<212> PRT

<213> group B streptococcus

<400> 27

Met Lys Asn Ile Thr Lys Leu Ser Thr Val Ala Leu Ser Leu Leu Leu			
1	5	10	15
Cys Thr Ala Cys Ala Ala Ser Asn Thr Ser Thr Ser Lys Thr Gln Ser			
20	25	30	
His His Pro Lys Gln Thr Lys Leu Thr Asp Lys Gln Lys Glu Glu Pro			
35	40	45	
Lys Asn Lys Glu Ala Ala Asp Gln Glu Met His Pro Gln Gly Ala Val			
50	55	60	

Asp Leu Thr Lys Tyr Lys Ala Lys Pro Val Lys Asp Tyr Gly Lys Lys
 65 70 75 80

Ile Asp Val Gly Asp Gly Lys Lys Met Asn Ile Tyr Glu Thr Gly Gln
 85 90 95

Gly Lys Ile Pro Ile Val Phe Ile Pro Gly Gln Ala Glu Ile Arg His
 100 105 110

Ala Met Leu Ile Arg Ile
 115

<210> 28

<211> 1191

<212> DNA

<213> group B streptococcus

<220>

<221> CDS

<222> (1)..(1191)

<400> 28

gtg aat gaa tcg acc atc aga aaa gaa ttt aaa ata gtt gtt ttt aaa 48
 Val Asn Glu Ser Thr Ile Arg Lys Glu Phe Lys Ile Val Val Phe Lys
 1 5 10 15

tgg atc tta aat aat caa gca gtt att gct ctc atg att acc ttt ttg 96
 Trp Ile Leu Asn Asn Gln Ala Val Ile Ala Leu Met Ile Thr Phe Leu
 20 25 30

gta ttt tta acg att ttt att ttt acc aaa atc tct ttt atg ttt aaa 144
 Val Phe Leu Thr Ile Phe Ile Phe Thr Lys Ile Ser Phe Met Phe Lys
 35 40 45

cct gtg ttt gat ttt ctt gct gtg ctg ata ttg ccg ctt gta att tct 192
 Pro Val Phe Asp Phe Leu Ala Val Leu Ile Leu Pro Leu Val Ile Ser
 50 55 60

ggc ttg ctt tat tac cta tta aaa cct atg gtt aca ttt tta gag aag 240
 Gly Leu Leu Tyr Tyr Leu Leu Lys Pro Met Val Thr Phe Leu Glu Lys
 65 70 75 80

cgg gga att aag cgt gta aca gcg ata tta tca gtt ttt act att ata 288
 Arg Gly Ile Lys Arg Val Thr Ala Ile Leu Ser Val Phe Thr Ile Ile
 85 90 95

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atc ctt ctg tta att tgg gca atg tct agt ttt att ccc atg atg agt   336
Ile Leu Leu Leu Ile Trp Ala Met Ser Ser Phe Ile Pro Met Met Ser
      100              105              110

aat caa tta cgc cat ttt atg gaa gat ctc cct tca tat gtg aat aaa   384
Asn Gln Leu Arg His Phe Met Glu Asp Leu Pro Ser Tyr Val Asn Lys
      115              120              125

gtg caa atg gaa aca agt tcg ttt ata gat cac aac cct tgg tta aaa   432
Val Gln Met Glu Thr Ser Ser Phe Ile Asp His Asn Pro Trp Leu Lys
      130              135              140

tct tat aaa ggg gaa ata tcg agc atg tta tct aat atc agt agc caa   480
Ser Tyr Lys Gly Glu Ile Ser Ser Met Leu Ser Asn Ile Ser Ser Gln
      145              150              155              160

gcg gtc tct tat gct gaa aaa ttt tca aag aat gtt tta gat tgg gca   528
Ala Val Ser Tyr Ala Glu Lys Phe Ser Lys Asn Val Leu Asp Trp Ala
      165              170              175

gga aat tta gct agt aca gtt gca cgt gtg aca gta gca aca atc atg   576
Gly Asn Leu Ala Ser Thr Val Ala Arg Val Thr Val Ala Thr Ile Met
      180              185              190

gct ccc ttt att ttg ttt tat ctt tta aga gat agt cgc aac atg aag   624
Ala Pro Phe Ile Leu Phe Tyr Leu Leu Arg Asp Ser Arg Asn Met Lys
      195              200              205

aat ggt ttc tta atg gtt tta cca acc aaa cta cgc caa cca gct gat   672
Asn Gly Phe Leu Met Val Leu Pro Thr Lys Leu Arg Gln Pro Ala Asp
      210              215              220

cgt att ttg cga gaa atg aat agt caa atg tca gga tat gtg caa gga   720
Arg Ile Leu Arg Glu Met Asn Ser Gln Met Ser Gly Tyr Val Gln Gly
      225              230              235              240

caa atc att gtt gct att act gtt ggt gtt att ttt tca ata atg tat   768
Gln Ile Ile Val Ala Ile Thr Val Gly Val Ile Phe Ser Ile Met Tyr
      245              250              255

agt att ata ggc ctt aga tat ggc gtg aca tta ggg att att gcc ggt   816
Ser Ile Ile Gly Leu Arg Tyr Gly Val Thr Leu Gly Ile Ile Ala Gly
      260              265              270

gtg tta aat atg gtt ccc tat ttg gga agt ttt gtc gcc caa att cca   864
Val Leu Asn Met Val Pro Tyr Leu Gly Ser Phe Val Ala Gln Ile Pro
      275              280              285

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gtg ttt atc tta gcg ctt gtc gca gga cct gtt atg gtt gtt aaa gtt 912
 Val Phe Ile Leu Ala Leu Val Ala Gly Pro Val Met Val Val Lys Val
 290 295 300

gcg att gtt ttt gtt att gag caa act cta gag gga cgc ttt gtc tca 960
 Ala Ile Val Phe Val Ile Glu Gln Thr Leu Glu Gly Arg Phe Val Ser
 305 310 315 320

cct ttg gtt tta ggt aat aaa ctt agc att cat cca att aca att atg 1008
 Pro Leu Val Leu Gly Asn Lys Leu Ser Ile His Pro Ile Thr Ile Met
 325 330 335

ttt att tta tta acc tct gga gcg atg ttt ggt gtt tgg gga gta ttc 1056
 Phe Ile Leu Leu Thr Ser Gly Ala Met Phe Gly Val Trp Gly Val Phe
 340 345 350

ctc agt att ccg att tat gca tct atc aaa gtt gtt gtt aaa gaa ttg 1104
 Leu Ser Ile Pro Ile Tyr Ala Ser Ile Lys Val Val Val Lys Glu Leu
 355 360 365

ttt gat tgg tac aaa gct gtc agt ggg cta tat aca ata gat gtt gtt 1152
 Phe Asp Trp Tyr Lys Ala Val Ser Gly Leu Tyr Thr Ile Asp Val Val
 370 375 380

act gaa gaa aga agt gaa gaa gtt aaa aat gtt gaa tag 1191
 Thr Glu Glu Arg Ser Glu Glu Val Lys Asn Val Glu
 385 390 395

<210> 29

<211> 396

<212> PRT

<213> group B streptococcus

<400> 29

Val Asn Glu Ser Thr Ile Arg Lys Glu Phe Lys Ile Val Val Phe Lys
 1 5 10 15

Trp Ile Leu Asn Asn Gln Ala Val Ile Ala Leu Met Ile Thr Phe Leu
 20 25 30

Val Phe Leu Thr Ile Phe Ile Phe Thr Lys Ile Ser Phe Met Phe Lys
 35 40 45

Pro Val Phe Asp Phe Leu Ala Val Leu Ile Leu Pro Leu Val Ile Ser
 50 55 60

Gly Leu Leu Tyr Tyr Leu Leu Lys Pro Met Val Thr Phe Leu Glu Lys
 65 70 75 80

Arg Gly Ile Lys Arg Val Thr Ala Ile Leu Ser Val Phe Thr Ile Ile
 85 90 95

Ile Leu Leu Leu Ile Trp Ala Met Ser Ser Phe Ile Pro Met Met Ser
 100 105 110

Asn Gln Leu Arg His Phe Met Glu Asp Leu Pro Ser Tyr Val Asn Lys
 115 120 125

Val Gln Met Glu Thr Ser Ser Phe Ile Asp His Asn Pro Trp Leu Lys
 130 135 140

Ser Tyr Lys Gly Glu Ile Ser Ser Met Leu Ser Asn Ile Ser Ser Gln
 145 150 155 160

Ala Val Ser Tyr Ala Glu Lys Phe Ser Lys Asn Val Leu Asp Trp Ala
 165 170 175

Gly Asn Leu Ala Ser Thr Val Ala Arg Val Thr Val Ala Thr Ile Met
 180 185 190

Ala Pro Phe Ile Leu Phe Tyr Leu Leu Arg Asp Ser Arg Asn Met Lys
 195 200 205

Asn Gly Phe Leu Met Val Leu Pro Thr Lys Leu Arg Gln Pro Ala Asp
 210 215 220

Arg Ile Leu Arg Glu Met Asn Ser Gln Met Ser Gly Tyr Val Gln Gly
 225 230 235 240

Gln Ile Ile Val Ala Ile Thr Val Gly Val Ile Phe Ser Ile Met Tyr
 245 250 255

Ser Ile Ile Gly Leu Arg Tyr Gly Val Thr Leu Gly Ile Ile Ala Gly
 260 265 270

Val Leu Asn Met Val Pro Tyr Leu Gly Ser Phe Val Ala Gln Ile Pro
 275 280 285

Val Phe Ile Leu Ala Leu Val Ala Gly Pro Val Met Val Val Lys Val
 290 295 300

Ala Ile Val Phe Val Ile Glu Gln Thr Leu Glu Gly Arg Phe Val Ser
 305 310 315 320

Pro Leu Val Leu Gly Asn Lys Leu Ser Ile His Pro Ile Thr Ile Met
 325 330 335

Phe Ile Leu Leu Thr Ser Gly Ala Met Phe Gly Val Trp Gly Val Phe
 340 345 350

Leu Ser Ile Pro Ile Tyr Ala Ser Ile Lys Val Val Val Lys Glu Leu
 355 360 365

Phe Asp Trp Tyr Lys Ala Val Ser Gly Leu Tyr Thr Ile Asp Val Val
 370 375 380

Thr Glu Glu Arg Ser Glu Glu Val Lys Asn Val Glu
 385 390 395

<210> 30

<211> 1230

<212> DNA

<213> group B streptococcus

<220>

<221> CDS

<222> (1)..(1230)

<400> 30

atg ttt atg gga atc cca caa tat ttc ttc tac ctt atc tta gct gtc 48

Met Phe Met Gly Ile Pro Gln Tyr Phe Phe Tyr Leu Ile Leu Ala Val

1 5 10 15

cta cca att tac atc ggc tta ttc ttt aag aag cgt ttt gcc tta tat 96

Leu Pro Ile Tyr Ile Gly Leu Phe Phe Lys Lys Arg Phe Ala Leu Tyr

20 25 30

gag att att ttt agt cta agt ttt att gta atg atg ttg act ggt agt 144

Glu Ile Ile Phe Ser Leu Ser Phe Ile Val Met Met Leu Thr Gly Ser

35 40 45

act ttt aat caa ttg aag tca cta ttg gca tac gtt gtc gga cag tct 192

Thr Phe Asn Gln Leu Lys Ser Leu Leu Ala Tyr Val Val Gly Gln Ser

50 55 60

ctg cta gtt ttt atc tat aaa gct tac cgg aaa cga ttt aat cat act 240

Leu Leu Val Phe Ile Tyr Lys Ala Tyr Arg Lys Arg Phe Asn His Thr

65 70 75 80

ttg gtc ttt tat gta acg gtt tgt tta tct att ttt ccg cta ttt ttg 288

Leu Val Phe Tyr Val Thr Val Cys Leu Ser Ile Phe Pro Leu Phe Leu
 85 90 95

gta aaa tta att cca gct ata tct gag gat ggg cat cag tca ctt ttt 336
 Val Lys Leu Ile Pro Ala Ile Ser Glu Asp Gly His Gln Ser Leu Phe
 100 105 110

ggg ttt cta gga att tct tac ctt act ttt aga gct gtt gct atg att 384
 Gly Phe Leu Gly Ile Ser Tyr Leu Thr Phe Arg Ala Val Ala Met Ile
 115 120 125

att gaa atg aga gac ggt gtc ttg aaa gaa ttt act tta tgg gaa ttc 432
 Ile Glu Met Arg Asp Gly Val Leu Lys Glu Phe Thr Leu Trp Glu Phe
 130 135 140

tta aga ttt tta ctc ttc ttt cca act ttc tca agt gga cca att gat 480
 Leu Arg Phe Leu Leu Phe Phe Pro Thr Phe Ser Ser Gly Pro Ile Asp
 145 150 155 160

cgt ttt aaa cga ttc aat gag gat tac att aat atc cca gat cga aac 528
 Arg Phe Lys Arg Phe Asn Glu Asp Tyr Ile Asn Ile Pro Asp Arg Asn
 165 170 175

gaa ctc cta gat atg tta ggt caa gcg att cat tat ttg atg tta ggt 576
 Glu Leu Leu Asp Met Leu Gly Gln Ala Ile His Tyr Leu Met Leu Gly
 180 185 190

ttt ctc tat aag ttt att tta gcc tat att ttt gga agt ctg att atg 624
 Phe Leu Tyr Lys Phe Ile Leu Ala Tyr Ile Phe Gly Ser Leu Ile Met
 195 200 205

cct cct cta aaa gaa tta gcg cta gaa cag ggt ggt gtg ttt aat tgg 672
 Pro Pro Leu Lys Glu Leu Ala Leu Glu Gln Gly Gly Val Phe Asn Trp
 210 215 220

cca aca ctt ggg gtt atg tat gcc ttt ggt ttt gat ttg ttc ttt gat 720
 Pro Thr Leu Gly Val Met Tyr Ala Phe Gly Phe Asp Leu Phe Phe Asp
 225 230 235 240

ttt gca ggt tac aca atg ttt gcg ttg gct att tct aac cta atg ggg 768
 Phe Ala Gly Tyr Thr Met Phe Ala Leu Ala Ile Ser Asn Leu Met Gly
 245 250 255

att aag tct ccg att aac ttt gac aaa cct ttc aaa tca cgc gac cta 816
 Ile Lys Ser Pro Ile Asn Phe Asp Lys Pro Phe Lys Ser Arg Asp Leu
 260 265 270

aaa gaa ttt tgg aat aga tgg cat atg agc ctt tct ttc tgg ttt aga 864

Lys Glu Phe Trp Asn Arg Trp His Met Ser Leu Ser Phe Trp Phe Arg
 275 280 285
 gac ttt gtt ttc atg agg ctt gtt aag ctt tta gtt aaa aat aaa gtt 912
 Asp Phe Val Phe Met Arg Leu Val Lys Leu Leu Val Lys Asn Lys Val
 290 295 300
 ttt aaa aac cgt aat gtt act tca agt gta gct tat att atc aat atg 960
 Phe Lys Asn Arg Asn Val Thr Ser Ser Val Ala Tyr Ile Ile Asn Met
 305 310 315 320
 ctt ctt atg gga ttc tgg cat ggg tta act tgg tac tat ata gcc tat 1008
 Leu Leu Met Gly Phe Trp His Gly Leu Thr Trp Tyr Tyr Ile Ala Tyr
 325 330 335
 ggt ctc ttt cat ggg att ggc cta gtt att aat gac gct tgg gta cgt 1056
 Gly Leu Phe His Gly Ile Gly Leu Val Ile Asn Asp Ala Trp Val Arg
 340 345 350
 aag aag aaa aat ayt aat aaa gaa aga aga ttg gct aaa aaa cca ctt 1104
 Lys Lys Lys Asn Xaa Asn Lys Glu Arg Arg Leu Ala Lys Lys Pro Leu
 355 360 365
 tta cca gaa aac aaa tgg act tat gct ttg ggt gtc ttc atc acc ttt 1152
 Leu Pro Glu Asn Lys Trp Thr Tyr Ala Leu Gly Val Phe Ile Thr Phe
 370 375 380
 aat gta gtt atg ttt tct ttc ttg att ttt tca gga ttt tta gat ctt 1200
 Asn Val Val Met Phe Ser Phe Leu Ile Phe Ser Gly Phe Leu Asp Leu
 385 390 395 400
 ttg tgg ttc cca caa ccg cat aac aaa taa 1230
 Leu Trp Phe Pro Gln Pro His Asn Lys
 405 410

<210> 31

<211> 409

<212> PRT

<213> group B streptococcus

<400> 31

Met Phe Met Gly Ile Pro Gln Tyr Phe Phe Tyr Leu Ile Leu Ala Val
 1 5 10 15

Leu Pro Ile Tyr Ile Gly Leu Phe Phe Lys Lys Arg Phe Ala Leu Tyr
 20 25 30

Glu Ile Ile Phe Ser Leu Ser Phe Ile Val Met Met Leu Thr Gly Ser
 35 40 45

Thr Phe Asn Gln Leu Lys Ser Leu Leu Ala Tyr Val Val Gly Gln Ser
 50 55 60

Leu Leu Val Phe Ile Tyr Lys Ala Tyr Arg Lys Arg Phe Asn His Thr
 65 70 75 80

Leu Val Phe Tyr Val Thr Val Cys Leu Ser Ile Phe Pro Leu Phe Leu
 85 90 95

Val Lys Leu Ile Pro Ala Ile Ser Glu Asp Gly His Gln Ser Leu Phe
 100 105 110

Gly Phe Leu Gly Ile Ser Tyr Leu Thr Phe Arg Ala Val Ala Met Ile
 115 120 125

Ile Glu Met Arg Asp Gly Val Leu Lys Glu Phe Thr Leu Trp Glu Phe
 130 135 140

Leu Arg Phe Leu Leu Phe Phe Pro Thr Phe Ser Ser Gly Pro Ile Asp
 145 150 155 160

Arg Phe Lys Arg Phe Asn Glu Asp Tyr Ile Asn Ile Pro Asp Arg Asn
 165 170 175

Glu Leu Leu Asp Met Leu Gly Gln Ala Ile His Tyr Leu Met Leu Gly
 180 185 190

Phe Leu Tyr Lys Phe Ile Leu Ala Tyr Ile Phe Gly Ser Leu Ile Met
 195 200 205

Pro Pro Leu Lys Glu Leu Ala Leu Glu Gln Gly Gly Val Phe Asn Trp
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Pro Thr Leu Gly Val Met Tyr Ala Phe Gly Phe Asp Leu Phe Phe Asp
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Phe Ala Gly Tyr Thr Met Phe Ala Leu Ala Ile Ser Asn Leu Met Gly
 245 250 255

Ile Lys Ser Pro Ile Asn Phe Asp Lys Pro Phe Lys Ser Arg Asp Leu
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Lys Glu Phe Trp Asn Arg Trp His Met Ser Leu Ser Phe Trp Phe Arg
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Asp Phe Val Phe Met Arg Leu Val Lys Leu Leu Val Lys Asn Lys Val
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Phe Lys Asn Arg Asn Val Thr Ser Ser Val Ala Tyr Ile Ile Asn Met
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Leu Leu Met Gly Phe Trp His Gly Leu Thr Trp Tyr Tyr Ile Ala Tyr
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Gly Leu Phe His Gly Ile Gly Leu Val Ile Asn Asp Ala Trp Val Arg
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Lys Lys Lys Asn Xaa Asn Lys Glu Arg Arg Leu Ala Lys Lys Pro Leu
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Leu Pro Glu Asn Lys Trp Thr Tyr Ala Leu Gly Val Phe Ile Thr Phe
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tgc gtt gga tgt tct gcc aat aaa gat aat caa aaa act aaa act gag 96
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 Glu Tyr Gly Leu Gly Val Ser Thr Gly Leu Asp Leu Pro Glu Ser Glu
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 ggt tat gta cct gga aaa tac agc tta gga aca act cta atg gaa tcg 144
 Gly Tyr Val Pro Gly Lys Tyr Ser Leu Gly Thr Thr Leu Met Glu Ser
 35 40 45
 ttc ggt cag tat gat gcc tat aca cca atg caa ctt ggt cag tat atc 192
 Phe Gly Gln Tyr Asp Ala Tyr Thr Pro Met Gln Leu Gly Gln Tyr Ile
 50 55 60
 tca act att gcg aat aat ggg aat cgt tta gca cct cac gtg gtt tca 240
 Ser Thr Ile Ala Asn Asn Gly Asn Arg Leu Ala Pro His Val Val Ser
 65 70 75 80
 gat atc tat gaa ggg aat gat tct aat aag ttc gct caa ttg gtt cgt 288

Asp Ile Tyr Glu Gly Asn Asp Ser Asn Lys Phe Ala Gln Leu Val Arg
 85 90 95

 tca atc act cct aaa aca cta aat aag ata gct atc tca gat caa gag 336
 Ser Ile Thr Pro Lys Thr Leu Asn Lys Ile Ala Ile Ser Asp Gln Glu
 100 105 110

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 Leu Ala Ile Ile Gln Glu Gly Phe Tyr Asn Val Val Asn Ser Gly Ser
 115 120 125

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 Gly Tyr Ala Thr Gly Thr Ser Met Arg Gly Asn Val Thr Thr Ile Xaa
 130 135 140

 ggt aaa act ggt acc gct gaa aca ttt gct aaa aat ata aat gga caa 480
 Gly Lys Thr Gly Thr Ala Glu Thr Phe Ala Lys Asn Ile Asn Gly Gln
 145 150 155 160

 aca gtt tct acc tac aac tta aac gct att gcc tac gat act aat cgt 528
 Thr Val Ser Thr Tyr Asn Leu Asn Ala Ile Ala Tyr Asp Thr Asn Arg
 165 170 175

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 Lys Ile Ala Val Ala Val Met Tyr Pro His Val Thr Thr Asp Thr Thr
 180 185 190

 aaa tcc cat caa tta gtt gca cga gat atg att gat caa tat att tca 624
 Lys Ser His Gln Leu Val Ala Arg Asp Met Ile Asp Gln Tyr Ile Ser
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 35 40 45

Phe Gly Gln Tyr Asp Ala Tyr Thr Pro Met Gln Leu Gly Gln Tyr Ile
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Ser Thr Ile Ala Asn Asn Gly Asn Arg Leu Ala Pro His Val Val Ser
 65 70 75 80

Asp Ile Tyr Glu Gly Asn Asp Ser Asn Lys Phe Ala Gln Leu Val Arg
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Ser Ile Thr Pro Lys Thr Leu Asn Lys Ile Ala Ile Ser Asp Gln Glu
 100 105 110

Leu Ala Ile Ile Gln Glu Gly Phe Tyr Asn Val Val Asn Ser Gly Ser
 115 120 125

Gly Tyr Ala Thr Gly Thr Ser Met Arg Gly Asn Val Thr Thr Ile Xaa
 130 135 140

Gly Lys Thr Gly Thr Ala Glu Thr Phe Ala Lys Asn Ile Asn Gly Gln
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Thr Val Ser Thr Tyr Asn Leu Asn Ala Ile Ala Tyr Asp Thr Asn Arg
 165 170 175

Lys Ile Ala Val Ala Val Met Tyr Pro His Val Thr Thr Asp Thr Thr
 180 185 190

Lys Ser His Gln Leu Val Ala Arg Asp Met Ile Asp Gln Tyr Ile Ser
 195 200 205

Gln Ser Gln Asp Asn Lys Arg Gly His
 210 215